								4	60						
trg	Lys 290	Va.1	Leu	Glu	Leu	Ala Al 295	a Ala	Leu	Ser	App 300	Amp	Phe	Glu	Arg	
11a	Gly	Arg	Arg												
(2)	INFO	ORMA:	rion	FOR	880	ID NO:	95:								
	(i)	SEC	JUEN	CE C	IARA	CIBRIST	ICS:								
						34 base		S							
						cleic a									
						NESS: s									
			(D)	TOPOI	LOGY	: linea	X.								
	(xi)	SE	SOEM	CE DI	ESCR.	IPTION:	SEQ	ID NO	; 9	5:					
AAG	ACTAC	ar :	TAT	GATO	ac o	BAGGATG	TT CO	CS							34
(2)	INFO	ORMA	TION	FOR	SEQ	ID NO:	96:								
	143	SEC	VF 1003345	ce c	er to be	CTERIST	****								
						27 base		8							
			(8)	TYPE:	rate	cleic a	cid								
						W295: 8									
			(D)	TOPOI	POGA	: linea	*								
	(xi)	ar.	SORN	CE DI	SSCR:	IPTION:	SEQ	ID NO	c 9	6:					
000	CGAC	rac (GGRT	CCTAS	ec G	CGTCGG									27
(2)	INFO	orma:	FICE	FOR	SEQ	ID NO:	97:								
	(1)	SE	OUEN	es e	er pr	CTERLST	ICS:								
						28 base		6							
			(8)	TYPE	nu	cleic a	cia								
						RESS: S									
			(D)	TOPO	LOGY	: lines	к								
	(xi	5.8	OCHER	CE DI	ESCR	iption:	980	TO NO	: 9	7:					
ccr	TGGG	AGA '	rem	TGGA	00 O	coerrec									28
(2)	INFO	ema'	rion	FOR	SEQ	ID NO:	98:								
	(i)	SS	XIEN	CE C	IARA	CTERIST	ICS:								
						25 base		8							
						cleic a									
						NESS: 8									
			5773	THY YOUNG	LALAN.	liman	24"								

	206	
(xi) SEQUENCE DESCRIPTION:	SRQ ID NO: 98:	
GACGAGATOT TATGGGCTTA CTGAC		25
(2) INFORMATION FOR SEQ ID NO:	99:	
(1) SEQUENCE CHARACTERIST (A) LEMBIN: 33 base (B) TYPE: nucleic a (C) STRANDEDMESS: s (D) TOPOLOGY: linea.	pairs cid ingle	
(wi) SEQUENCE DESCRIPTION:	SEQ ID NO: 99:	
CCCCCCAGAT CTGCACCACC GGCATCGG	CG GGC	33
(2) INFORMATION FOR SEC ID NO:	100	
(i) SEQUENCE CHARACTERIST (A) LENGTH: 24 base (B) TYPE: nucleic a (C) STRANDEDNESS: s (D) TOPOLOGY: linea	pairs aid ingle	
(xi) DEQUENCE DESCRIPTION:	SEQ ID NG: 190:	
GCGGCGGATC COTTGCTTAG COGG		24
(2) INFORMATION FOR SEQ ID NO:	101:	
(i) SEQUENCE CHARACTERIST (A) LEMMIN: 32 base (B) TYPE: nucleic a (C) STEARDEDNESS: s (D) TOPOLOGY: linea	pairs cid ingle	
(xi) SEQUENCE DESCRIPTION:	SEQ ID 90: 161:	
OCCOCTGAGA TOTATGACAG AATACGAA	se ec	3.2
(2) INFORMATION FOR SEQ ID NO:	102 -	
(i) SEQUENCE CHARACTERIST (A) LENGTH: 24 base (B) TYPE; nuclaic a (C) STRANDEDMESS: s (D) TOPOLOGY: linea	pairs cid ingle	
(x1) SEQUENCE DESCRIPTION:	SEQ ID NO: 102:	
CCCCGCCAGG GAACTAGAGG CGGC		24
(2) INFORMATION FOR SEC 1D NO:	103.	

(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 38 base pairs	
(B) TYPE: nucleic scid	
(C) STRANDEDNESS: single	
{D} TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 103:	
CTGCCGAGAT CTACCACCAT TGTCGCGCTG AAATACCC	38
(2) INFORMATION FOR SEQ ID NO: 164:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 25 base pairs	
(B) TYPE: mucleic exid	
(C) STRANDEDNESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 104:	
CGCCATGGCC TTACGCGCCA ACTCG	25
(2) IMPORMATION FOR SEQ ID NO: 105:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LEMGTH: 12 base pairs	
(B) TYPE: nucleic acid	
(C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(D) TOPCHOUS: IInear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 105:	
GGCOGAGATC TOTGAGTTTT CCGTATTTCA TC	32
(2) INFORMATION FOR SEQ ID NO: 106:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 25 base pairs	
(E) TYPE: mucleic acid	
(C) STWANDEDNESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 196:	
CGCGTCGAGC CATGGTTAGG CGCAG	25
(2) INFORMATION FOR SEQ ID NO: 167:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 32 base pairs	
(B) TYME: nucleic acid (C) STRANDEONESS: single	
(D) TODOLOGY: 1 Mark	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 107:	
GAGGAAGATC TATBACAACT TCACCCGACC CC	32
(2) INFORMATION FOR SEQ ID NO: 108:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 28 base pairs	
(B) TYPE: nucleic acid	
(C) STRANDEDRESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION; SEQ ID NO: 108;	
CATGAAGCCA TGGCCCGCAG GCTGCATG	28

(2) INFORMATION FOR SEQ ID NO: 109:	
(i) sequence characteristics:	
(A) LENGTH: 33 base pairs	
(S) TYPE: nucleic acid	
(C) STRANDEDNESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 109:	
SUCCEMBATE TOWNSCREAC TATURCUTES TES	33
(2) IMPORMATION FOR SEQ ID NO: 110:	
(i) SECURNCE CHARACTERISTICS;	
(A) LENGTH: 36 base pairs	
(B) TYP8: nucleic acid	
(C) STRANDEDNESS: single	
(D) TOPOLOGY: linear	
(will SEQUENCE DESCRIPTION: SEQ ID NO: 116:	
7	
OGCGCCCATG GICAGAAATT GATCATGTGG CCAACC	36
(2) INFORMATION FOR SEQ ID NO: 111:	
(%) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 33 base pairs	
(B) TYPE: nucleic soid	
(C) STRANDEDRESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 111:	
COGGRATT TATGGCAAAG CTCTCCACCG ACG	33
(2) INFORMATION FOR SEQ ID NO: 112:	

	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 32 base pairs	
	(B) TYPE: aucleic acid	
	(C) STRANDEDMESS: single	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION; SEQ ID NO: 112:	
CGC	TEGECAG ASCTACTIGA CGETGACGET GG	32
(2)	INFORMATION FOR SEQ ID NO: 113:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LEWSTH: 36 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: single (D) TOPOLOGY: linear	
	(b) TOPOLASI: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 113:	
000	CCAGATC TATOSCCATT GAGGTTTCGG DETTOC	3.6
(2)	INFORMATION FOR SEQ ID NO: 114:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 26 base pairs	
	(B) TYPE: nucleic soid	
	(C) STRAMDEDRESS: single	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 114:	
CGC	COTOTTO CATOCCACCO CTGAGC	26
(2)	INFORMATION FOR SEQ ID NO: 115:	
	(1) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 24 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: single	
	(D) TOPOLOGY; linear	
	(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 125:	
GGA	ACCITICAR GOGACACRIC GOOG	24
(0)	Valentia Maria Dan Anna Anna Anna Anna Anna Anna Anna	
(2)	INFORMATION FOR SEQ ID NO: 116:	
	(1) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 24 base pairs (B) TYPS: nucleic scid	
	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	

	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 116:	
CAG	SCACGAAC GCGCCGTCGA 798C	24
(2)	INFORMATION FOR SEQ ID NO: 117:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LEMOTH: 26 base pairs	
	(h) TYPE: nucleic acid	
	(C) STRANDEDWSSS: single	
	(D) TOPOLOGY: linear	
	4.2) 4000000000000000000000000000000000000	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 117:	
ACE	AGRICITY GACGGACATG AACCCG	26
(2)	INFORMATION FOR SMQ ID NO: 118:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENJTH: 28 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(*i) SEQUENCE DESCRIPTION: SEQ ID NO: 118:	
TT	TTCCATOG TCACGGGCCC COXGTACT	28
(23	I IMPORMATION FOR SEQ ID NO: 119:	
	(i) SECURECE CHARACTERISTICS:	
	(A) LENGTH: 36 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDHDNRSS single	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 115:	
AC	agatetut gecertoger eagata	26
(2)) INFORMATION FOR SEQ ID NO: 128;	
	, and the same and	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LEMSTH: 27 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDNESS: mingle	
	(D) TOPOLOGY: linear	
	7-11 AMONOMONE PRESIDENTAL - \$100 10 MO. 100.	
	(xi) SEQUENCE DESCRIPTION: SEQ 1D NO: 120:	
TT	TAASCTIC TAGGODOCCA SCGCGGC	27
12	INFORMATION FOR SEQ ID NO: 121:	

211

(i) REPRIENCE CHARACTERICO.

(A) LEMOTH: 26 base pairs (B) TYPE: mucleic acid (C) STRAMBERNESS: single (D) TOPOLOSY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO; 121:	
ACAGATETEC GCATGCOGAT COGTGT	2
(2) INFORMATION FOR SEQ ID NO: 122:	
(i) SEQUENCE CHARACTERISTICS: (A) LEMSTH: 26 base pairs (B) TYPE: mucleic acid (C) STRAMBEDNESS: single (D) TOPOLOGY: linear	
(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 122:	
TTTTCCATGG TCATCCGGCG TGATCGAG	2
(2) INFORMATION FOR SEQ ID NO: 123;	
(i) SEQUENCE CHARACTERISTICS: (A) LEMOTH: 25 have pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION; SEQ ID NO: 123:	
ACMUNTCTOT ANTOGORGAC TOTOAT	2
(2) INFORMATION FOR SEQ ID NO: 124:	
(i) SEQUENCE CHARACTERISTICS: (A) LEMOTH: 28 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(wi) SEQUENCE DESCRIPTION: SEQ ID NO: 124:	
TTTTCCATCG TCACCAGATG GTGATCGA	Ž
(2) IMPORMATION FOR SEQ ID NO: 125:	
(1) SEQUENCE CHARACTERISTICS: (A) LEMOTH; 26 base pairs (B) TYPE: Nucleic acid (C) STRANDENNESS: single (D) TOMOGRAPHICAL	

212 (xi) SECURNCE DESCRIPTION: SEC ID NO: 125: ACAGATOTGC CGGCTACCCC GGTGCC 26 (2) INFORMATION FOR SEQ ID NO: 126: (i) SEQUENCE CHARACTERISTICS: (A) LEWSTH: 28 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (mi) SEQUENCE DESCRIPTION: SEQ ID NO: 126: TTTTCCATGG CTATTGCAGC TTTCCGGC 28 (2) INFORMATION FOR SEQ ID NO: 127: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 50 amino acids (B) TYPE: amino acid (C) STRANDEONESS; single (D) TOPOLOGY: linear (iii) MOLECULE TYPE: None (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 127: Ala Glu Asp Val Arg Ala Glu Ila Val Ala Ser Val Leu Glu Val Val Val Asn Glu Gly Asp Sln Ile Asp Lys Sly Asp Val Val Val Leu Leu 25 Glu Sex Met Tyr Met Glu Ile Pro Val Leu Ala Glu Ala Ala Gly Thr 4.0 Val Ser 50

- (2) INFORMATION FOR SEQ ID NO: 128:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LEMFTH: 49 amino acida
 - (B) TYPE: amino acid
 - (C) STRANDEDWESS: single
 - (D) TOPOLOGY: linear
 - (ii) MOLECULE TYPE: None
 - (xi) SMOUTENCE DESCRIPTION: SEO ID NO. 128:

Ala Glu Asp Val Arg Ala Glu Ile Val Ala Ser Val Leu Glu Val Val 1 5

Val Asn Glu Gly Asp Gln Tie Asp Lys Gly Asp Val Val Leu Leu 20 25 3.0

213 Glu Ser Met Met Glu Ile Pro Val Leu Ala Glu Ala Ala Gly Thr Val 35 40 Ser (2) INFORMATION FOR SEQ ID NO: 129: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 50 amino acids (B) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: None (xi) SECURNCE DESCRIPTION: SEC ID NO: 129: Ala Glu Asp Val Arq Ala Glu Ile Val Ala Ser Val Leu Glu Val Val 3 Val Acn Glu Gly Asp Gln Ile Asp Lys Gly Asp Val Val Val Leu Leu 25 Glu Ser Met Lys Met Glu Ile Pro Val Leu Ala Glu Ala Ala Gly Thr 35 40 Val Ser 50 (2) INFORMATION FOR SEC ID NO: 130: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 33 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (b) TOPOLOGY: linear (xi) SHOURNCE DESCRIPTION: SEC 10 NO: 130: COGGGAGATO TATOGCAAAG CTCTCCACCG ACG 33 (2) INFORMATION FOR SEQ ID NO: 131: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (xi) SECUENCE DESCRIPTION: SRO ID NO: 131: CGCTGGGCAG AGCTACTTGA CGGTGACGGT GG 32

(2) INFORMATION FOR SHO ID NO: 132:

(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 36 base pairs	
(B) TYPB; nucleic acid	
(C) STRANDEDNESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ 1D NO: 132:	
, , , , , , , , , , , , , , , , , , ,	
OUCUCCUUCA AUCTIUCCAT GACAGAGCAG CAGTOG	36
(2) INFORMATION FOR SEQ ID NO: 133;	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 26 base pairs	
(B) TYPE: nucleic acid	
(C) STRAMDEDNESS: single	
(D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 183:	
(see) pageance seems to be an en. 200.	
CGAACTOGCC GGATCCCOTG TTTCGC	26
(2) INFORMATION FOR SEQ ID NO: 134:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 32 base pairs	
(B) TYPE: nucleic acid	
(C) STRANDEDNESS: single	
(D) TOPOLOGY: linear	
(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 134:	
	32
GGCRACCGCG AGATCTTTCT CCCGGCCGGG GC	34
(2) INFORMATION FOR SEQ ID NO: 135:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 27 base pairs	
(%) TYPE: nucleic acid	
(C) STRANDEDNESS: single (D) TOPOLOGY: linear	
in invitant inext	
7.27	
(xi) SECREBCE DESCRIPTION, SEC ID NO: 135;	
GGCAAGCTT% CORECGCCTA ACGAACT	27
ALL MINISTERS AND ADDRESS AND	
(2) INFORMATION FOR SEQ ID NO: 136:	
(i) SEQUENCE CHARACTERISTICS:	
(A) LENGTH: 36 base pairs	
(B) TYPE: nucleic acid	
(C) STRANDEDNESS: single	
(D) Which hay, linear	

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 136:	
GGACCCAGAT CTATGACAGA GCAGCAGTGG	30
(2) INFORMATION FOR SEQ ID NO: 137:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 47 base pairs (B) TYPE: Rucloid acid (C) STRANDEDNESS: Single (D) TOPOLOGY: linear	
(x1) SEQUENCE DESCRIPTION: SEQ ID NO: 137:	
CCGGCAGCCC CGGCCGGGAG AAAAGCTTTTG CGAACATCCC AGTWACG	-67
(2) INFORMATION FOR SEQ ID NO: 138:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 44 base pairs (B) TYPE: nucleic acid (C) STRANDENNESS: single (D) TOPOLOGY: linear	
(%i) SEQUENCE DESCRIPTION; SEQ ID NO: 138:	
STTCSCAAAS CTTTTCTCCC SSCCSSSSCT GCCGSTCSAG TACC	44
(2) INFORMATION FOR SEQ ID NO: 139:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 26 base pairs (B) TTPR: next ein acid (C) STRANDEDNESS: single (D) TOPCSOSY: linear	
(x1) SEQUENCE DESCRIPTION; SEQ ID NO: 139:	
CCTTCGGTGG ATCCCGTCAG	26
(2) IMPÓRMATION FOR SEQ IS NO: 140:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 450 base pairs (B) TIPE: mucleic acid (C) STRANGENESS: single (D) TOPOLOGY: linear	
(ix) FRATURE:	
(A) NAME/KEY: Coding Sequence (B) LOCATION: 68346 (D) OTHER INFORMATION:	

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enc.	,							ero i								
-	CGCIA	326 3	rcca:	GGA	AC C	GTC	WTG	r car	rcgac	3CAG	TAC	rgaa	cca '	TCO	AAADAS	60
GC	CAGC															109
		Ment:	Asn	Val	Thx	Val	Ser	Tle	Pro	The	Ile 10	Leu	Arg	Pro	His	
		٠,١.				20					**					
	GGC															157
ar 25	Gly	Gly	Gln	Lys	Ser 20	Val	Ser	Ala	Ser	53y	Asp	3.17%	767	GIY	ALS 30	
43					20					#2					an ex	
	ATC															205
al	Zle	Ser	Asp	Leu 35	Glu	Ala	Asu	Tyr	Ser 40	Giy	Tle	Ser	Glu	Arg	Leu	
				33					40					œ.,		
	GAC															253
See 2	yaz	Pro	Ser	Ser	Pro	Gly	Lys	Les 55	His	Axg	Pfre	val	Aso 60	Ile	Tyx	
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al	Asm		Glu	Asp	Val	Arg		Ser	Gly	ggy	Leu		Thr	Ala	Tle	
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cr	GAC	GGT	GAC	300	GTC	ACC	ATC	CZC	CCC	GCC	GTG	GCC	GGT	GGG	TGAGC	351
(la	Asp	Giy	Asp	Ser	Val		He	Leu	Pro	Ala		Ala	Gly	gly		
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GA	GCAC.	ora	ACAC!	BATA	CG A	CTCG	CERET	rec	AGGCC	CTTG	GGC	AACA	COC 4	cecu	POTTEG	411
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(2)	INF	ORMA	TION	FOR	SBÖ	ID :	: : 0%	141:								
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	3.8															

Gly Gin Lys Ser Val Ser Ala Ser Gly Asp Thr Leu Gly Ale Val Ile Ser Asp Leu Glu Ala Asn Tyr Ser Cly Ila Ser Glo Arg Leu Mat Asp

Pro Ser Ser Pro Gly Lyg Leu His Arg Dhe Val Asn Ile Tyr Val Asn

60

55

35

									2	17						
Asp 65	Glu	Asp	Val.	Arg	Phe 70	Ser	Gly	Gly	Leti	Ala 75	Thr	Ala	The	Ala	Asp 80	
gly	Asp	Ser	Val	The 85	114	Leu	Pro	Ala	Val 90	Ala	Gly	Gly				
(2)	INFO	TRMA'	rion	FOR	SRQ	ro i	(O: 1	42:								
	(i)		(A) 1 (B) 1 (C) 1	CE CH LENGT TYPE: TRAN TOPOI	(B): 4 : nuc :DED	180 k Gleit NESS	case caci	pai:	rB							
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rec	AGGT	rca :	ACAM	IDAGE	AC AC	7GCA									AT COS sp Pro	114
	Ala								TTT							162
									GCG Ala 35							216
									GCG Ala							258
									ATC							306
		Asp							AAC							354
	Ala			CAG Cln					TAA	corc	AGC	cecr	CAG	CA C	AATACT	408
Selection of the select	ACAA	aca .	aagg.	AGAR	CA G	GTTO	eate	a cc	atca	acta	TCA	GTTC	GGT -	GATG	TOGACG	468
CTC	ATGG	cac	A)													480

218

(mi) SEQUENCE DESCRIPTION: SEC ID NO: 143:

Met Als Thr Arg Phe Met Thr Asp Pro His Als Met Arg Asp Met Als 1 18 15

Gly Arg Vhe Glu Vel His Ala Gln Thr Val Glu Aep Glu Ala Arg Arg
25
30
Met Trp Ala Ser Ala Gln Asn Ile Ser Gly Ala Gly Trp Ser Gly Met
35
45
Ala Glu Ala Thr Ser Leu Asp Thr Met Ala Gln Met Asm Gln Ala Phe
50
55

Arg Asm lie Val Asm Met Leu His Gly Val Arg Asm Gly Leu Val Arg 65 70 75 80 Asm Ala Asm Asm Tyr Glu Glm Glu Glu Ala Ser Glm Glm lie Leu 85 90 95

(2) INFORMATION FOR SEQ ID NO: 143;

(i) SEQUENCE CHARACTERISTICS:
(A) LENGTH: 98 amino acids
(B) TYPE: maino acid
(C) STEANDEDNESS: single
(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: protein
(v) PRAGMENT TYPE: internal

(2) INFORMATION FOR SEQ ID NO: 144:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 940 base pairs

(B) TYPE: nucleic acid

(C) STRANDEDMESS: single

Ser Ser

(D)	TOPOLOGY: 1	Linear					
(ix) FEATU	RS:						
(A)	NAME/KEY: (	Coding :	Sequenc	9			
(2)	LOCATION: 8	36 86	8				
(D)	OTHER INFO	MATION					
SCCCUAGTEC TOS	NCE DESCRIPT				CGACCGC	AGGCCAC	GTG 60
TOCGCCACCT AAC	GAAAGGA TGA:	rc arg	CCC ANG	AGA AGC	GAA TAG	AGG CA	A 112
		Mee	Pro Lys	Arg Ser	Glo Tys	Arg Gl	ž)
		1					
GGC ACG CCG AA	a man ama m	or come	020 200	300 000	conce more	000 00	2 160

									2	19						
Gly 10	The	Pro	Asn	Trp	Val 15	Asp	Leu	Gln	Thr	Thr 20	Asp	ala	Ser	Ala	Ala 25	
	aag Lys															208
	CCC Pro	Gly													GAA Glu	256
			Ala												ATG Met	384
	OCG Pro														GTG Val	352
	GAC Asp															400
	ATC															448
	GCC Ala			Len					Arg							496
			Thr												AAG Uys	544
	GAT Asp 155	Leu														592
	Sex														OCC Ala 185	640
	GAC				Gly											688
	AAT Asn			Nis											ACG Thr	735
			Ala					Gly							oct Ala	784
Asp	ATT 11e 235	Pro					Phe					Asp				832

220 GOG ATC TTC AGT GTG TTG AAG CCC GCA CCG CNG CAA TAGGGAAGCAT CCCGGC 884 Ala Ile Phe Ser Val Leu Lys Pro Ala Pro Gin Gin 255 CAGGCCCGCC GGCCGGCAGA TTCGGAGAAT GCTAGAAGCT GCCGCCGGCG CCGCCG 940 (2) INFORMATION FOR SEQ ID NO: 145: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 261 amino acids (B) TYPE: amino acid (C) STRANDEDNESS; single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (v) FRAGMENT TYPE: internal (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 145; Met Pro Lys Arg Ser Glu Tyr Arg Gln Gly Thr Pro Asn Trp Val Asp Leu Glo Thr Thr Asp Slo Ser Ale Ale Lys Lys Phe Tyr Thr Ser Leu 25 20 Phe Gly Trp Gly Tyr Asp Asp Asp Pro Val Pro Gly Gly Gly Giv Val Tyr Ser Met Ala Thr Leu Ash Gly Glu Ala Val Ala Ala Ile Ala Pro Met Pro Pro Gly Ala Pro Glu Gly Met Pro Pro Ile Trp Asn Thr Tyr 70 65 Ile Ala Val Asp Asp Val Asp Ala Val Val Asp Lys Val Val Pro Cly 9.0 Gly Gly Gln Val Met Met Pro Ala Phe Asp Ile Gly Asp Ala Gly Arg Met Ser Phe Ile Thr Asp Pro Thr Gly Ala Ala Val Gly Leu Trp Gla 335 1.20 Ala Asn Arg His Ile Gly Als Thr Leu Val Asn Glu Thr Gly Thr Leu 135 The Trp Asn Glo Leu Leu Thr Asp Lys Fro Asp Leu Ala Leu Ala Phe 345 150 155 Tyr Glu Als Val Val Cly Leu Thr His Ser Ser Met Clu Ile Ala Als 170 Gly Gln Asn Tyr Arg Val Leu Lys Ala Gly Asn Ala Glu Val Gly Gly

Cys Met Glu Pro Pro Mer Pro Gly Val Pro Asn His Trp His Val Tyr

205

300

221

Phe	Ala 210	Val	Asp	Asp	Ala	Asp 215	Ala	The	Ala	Ala	Lys 220	Ala	Ala	Rla	Ala	
Gly 225	Gly	Gln	Val	Tle	Ala 230	Glu	Pro	Als	Asp	11e 235	Pro	Ser	Val	Gly	Arg 240	
Phe	Ala	Val	Leci	Ser 245	Asp	Pxo	Gln		Ala 250	lle	Phe	Sex	val	Leu 255	Lys	
Pro	Ala	Pro	Gln 266	Gln												
{2}	INF(	ORMA!	rion	FOR	SEQ	ID I	9O: :	146:								
			(A) 1 (B) 1 (C) 1 (D) 1	CE CE LENGT EYPE STRAI TOPOI	TH: : : BUS : BUS	200 ) Clei NESS	Dase C ac:	pai:	rs							
			(B)	NAME, LOCAT	PION	47	2	47	aonc:	>						
	(xí	) se	OURN	CIE DI	RSCR.	IPTI	ON: :	SEQ :	ID M	): 1s	\$ S :					
CCG	Larg	GCG 1	etga:	ACCG(	CA C	CCAG	AAGA.	a aai	ggari	AGAT	CGAV			eca (	CAG Sin	ŚŚ
GGA	ACT	oro	aag	Tog	TTC	AAC	GCU	gag	AAG	oog	TTC	99C	Met. 1 1 TTT	Pro (	3in GCC	103
GGA Gly	ACT Thr 5	GTG Val	aag Lys GgT	TOG Trp	TTC Phe	AAC Asn 10	GCU Ala GTA	GAG Glu	AAC Lys	GGG Gly	TTC Phe 15	60C Gly	Met 1 1 TTT Phe	Pro (	GCC Ala CAG	
GGA Gly CCC Pro 20	ACT Thr 5 GAA Glu	GTG Val GAC Asp	AAG Lys GGT Gly	TGG Trp TCC Ser	TTC Phe GCG Ala 25	AAC Asn 10 GAT Asp	GCU Ala GTA Val	GAG Glu TTT The	AAC Lys GTC Val	GGG Gly CAC Ris 30	TTC Phe 15 TAC Tyt	69C Gly ACG Thr	Met I I TTT Phe GAG Glu GAG	ATC	GCC Ala CAG Gln 35	103
GGA Gly CCC Pro 20 30A Gly	ACT Thr 5 GAA Glu ACG Thr	GTG Val GAC Asp GGC Gly	AAG Lys GGT Gly TTC Phe	TOG Trp TCC Ser CGC Arg 40	TTC Phe GCG Ala 25 ACC Thx	AAC Asn 10 GAT Asp CTT Leu	GCU Ala GTA Val GAA GIU	GAG Glu TTT Vhe GAA Glu	AAC Lys GTC Val AAC Asn 45	GGG Gly CAC His 30 CAG Gln	TTC Phe 15 TAC Tyr AAG Lys	GOC Gly ACG Thr GTC Val	Met I I TTT Phe GAG Glu GAG Glu COC	ATC Ile ATC Ile TTC Phe 50	GCC Als CAG Gln 35 GAG Glu	103
GGA Gly CCC Pro 20 SGA Gly ATC Ile	ACT Thr 5 GAA Glu ACG Thr GGC Gly	GAC Asp GGC Gly CAC His	AAG Lys GGT Gly TTC Phe AGC Ser 55	TOG Trp TCC Ser CGC Arg 40	TTC Phe GCG Ala 25 ACC Th: AAG	AAC Asn 10 GAT Asp CTT Leu GGC Gly	GCU Als GTA Val GAA Glu CCC Pro	GAG Glu TTT Whe GAA Glu CAG Gln 60	AAC Lys GTC Val AAC Asn 45	GGG Gly CAC His 30 CAG Gln	TTC Phe 15 TAC Tyr AAG Lys	GOC Gly ACG Thr GTC Val	Met !  I TTT Phe GAG Glu GAG Glu CGC Arg	ATC Ile ATC Ile TTC Phe 50	GCC Als CAG Gln 35 GAG Glu	103 153 199
GGA Gly CCC Pro 20 SNA Gly ATC Ils	ACT The 5 GAA Glu ACG Thr GGC Gly	GAC Asp GGC Gly CAC His	ARG Lys GGT Gly TTC Phe AGC Ser 55	TOG Trp TCC Ser CGC Arg 40	TTC Phe GCG Ala 25 ACC Thx AAG Lys	AAC Asn 10 GAT Asp CTT Leu SGC Gly	OKU Ala GTA Val GAA Glu CCC Pro	GAG Glu TIT Phe GAA Glu CAG Gln 60	AAC Lys GTC Val AAC Asn 45	GGG Gly CAC His 30 CAG Gln	TTC Phe 15 TAC Tyr AAG Lys	GOC Gly ACG Thr GTC Val	Met !  I TTT Phe GAG Glu GAG Glu CGC Arg	ATC Ile ATC Ile TTC Phe 50	GCC Als CAG Gln 35 GAG Glu	103 153 199 248

{8} TYPE: amino soid
{C} STRANDEDNESS: single
{D} TOPOLOGY: linear

				ST T				i.								
	(xi.)	SE	QUEN	28 D8	sca	PTIC	M: 5	IRQ :	ID N	3; 16	7:					
Met 1	Pro	Gln	Gly	Thr 5	Val	Ъув	Trp	Phe	Asn 10	Ala	01u	Lys	Gly	Phe 15	Sly	
Phe	Ile	Ala	Pro 26		Asp	Gly	Ser	Ala 25	Asp	Val	Phe	Va1	His 30	Tyr	Thr	
ela	Ile	Gln 35	Gly	Thr	Gly	Phe	Arg 40	Thr	Leu	Glu	Glu	A#0 45	Gla	Lys	Val	
			Ile			Ser 55	Pro	Lys	Gly	Pro	Gln 60	Als	Thr	Gly	Val	
Arg 65	Ser	Leu														
(2)	INF	DRMR'	LION	FOR	QEQ	ID I	90: :	148:								
	{ <b>x</b> i	) Fr.	(C) (C) (D) (A) (A) (A) (B) (C) (D) (C)	NAME LOCA DINE CE DI	IDEDI LOGY /KEY FION R IN	NESS : li: : Cox : lo: FORM	: si: Dear ding 5 ATIC:	Seq Seq 191 %:	uesc ID N	D: 1·		2860	ana :	n never	ITYATA	€3
												G AT	AA G	C CT	A CGG	116
													1			
									Leu						ATT Ile 20	154
															GAC Asp	312
									Ala					Gly	gat Asp	260
			Ala					\$1n							SCC Ala	306

223

									2	223						
															CGC Arg	356
24.4	70	sar y	uzy	961	Pile	75	2112	Ala	VOL	Mid	86	A.G.T.	104	nia	MIN	
	CAA															404
85	Gla	ary	1965	ser	90	wsp	SHE C	MIS	OTE	95	Phe	1877	Sec	116	100	
	TCG Ser															452
6.60	1360 K	4000 S		105	110	2007	w 68.V	AMOS C	110		- tx -t	PALEX	COCON	115	24010	
	CCG												TAG	3CGTV	SCG CG	503
mes (1	110	317.07	120	PIO	Map	gate to	2.1.0	125	Date	FIU	era	aux				
CT	CTA	3CC 1	ggro	ATO	AC G	GATC	GATO	G TG	eatg							540
(2)	INFO	DBMY,	TION	POR	SEQ	ID I	<b>\$0:</b>	149:								
	(V)	MC FR	(C) ; (D) ' LECU AGME QUES	ropo le t nt t	LOGY YPE: YPE:	pro	near tein erna		ID N	Ot 1	19:					
%et 1	Asn	Leu	Arg	Arg 5		Gla	Thr	Leu	The 10		Arg	Leu	Leu	Ala 15	Ala	
Ser	Ala	Gly	Ile 20		Ser	Ala	Ala	Ala 25	Phe	Ala	Ala	Pro	Ala 30	Gln	Ala	
Asn	Pro	Val 35		Asp	Ala	Pite	11a 40		Ala	Lou	neA	Aan 45	Ala	Gly	Val	
Asn	772 50	Gly	Asp	Pro	Val	Asp 55	Als	Lys	Ala	Leu	60 614		ser	Val	Cys	
Pro 65	He	Leu	Ala	Glu	Pro 70		Oly	Ser	Phe	Asn 75	"hr	Ala	Val	Ala	Ser 80	
Val	Val	Als	Arg	Ala 85		Gly	Met	Ser	Gln 90		Met	Ala	Gln	Thr 95		
Thr	Ser	lle	Ala 100		Ser	Met		Cys 105		Ser	Val	Met	Ala 110		Val	
Ala	Ser	Gly 115		Leu	Pro	Ala	Leu 120		Asp	Nes	Pro	01y 125		Pro	Gly	
Ser																

Ser

224

INFORMATION			

## (i) SEGUENCE CHARACTERISTICS:

- (A) LENGTH: 400 base pairs (B) TYPE: nucleic acid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (ix) PRATTER:

- (A) NAME/KEY: Coding Sequence
- (B) LOCATION: 25...354 (C) OTHER INFORMATION:

# (ix) FEATURE:

- (A) NAME/KEY: mat_peptide
- (B) LOCATION: 169.357

## (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 150:

ATAGTTTGGG	GAAGGTGTCC	ATTAA	ATG	AGG	CTG	TCO	TTO	ACC	GCA	220	AGC
			Met	Arg	Leu	Ser	Leu	Thr	Ala	Leu	Ser
			. 28			-25					-20

GCC	GGT	GTA	ggc	GCC.	GTG	GCA	RTG	TCG	TTG	ACC	GTC	GGG	GCC	QQQ	GTC	9.9
Ala	Gly	Val	Gly	Ala	Val	Ala	Met	Sor	Leu	Thr	Val	Gly	Ala	Gly	Val	
				-15					- 20					5		

53.

GCC	TCC	GCA	anr	000	ara	CAC	GCG	orc	ATT	AAC	ACC	ACC	TOC	AAT	TAC	167
Ala	Ser	Ala	Asp	Pro	Val	Asp	Ala	Val	rie	Asn	Thr	Thr	Cys	Asn	Tyx	
							101					2.25				

GGG	CAG	GTA	GTA	GCT	GCG	CTC	AAC	GCG	ACG	GAT	COB	GGG	GCT	GCC	GCA	192
Gly	Gin	Val	Val	BIA	Ala	Leu	Asn	Ala	Thr	Asp	pro	Gly	Ala	Ala	Ala	
	2.5					20					25					

CAG	TTC	AAC	GCC	TON	CCC	GTG	GCG	CAG	TCC	TAT	TTG	COC	ART	TTC	CAC	243
Gin	Phe	Ann	Ala	Sex	Pro	Val	Als	Gin	Sex	Tyx	Lev	Arg	Asn	Phe	Leu	
30					35					40					45	

GCC	GCA	CCG	CCA	CCT	CAG	CGC	GCT	GCC	ATG	GCC	GCG	CAA	TTG	CAA	GCT	293
Ala	Ala	Pro	Pro	Pro	Gln	Arg	Ala	Ala	Met	Ala	Ala	Gln	Leu	Gln	Ala	
				50					55					60		

GTG	cca	ggg	GCG	GCA	CRG	TAC	ATC	GGC	CTT	GTC	GNG	TCG	GTT	GCC	GGC	335
Val	Pro	Gly	Ala	Ala	Glo	Tyx	110	Gly	Leu	Val.	Glu	Ser	Val	Ala	Gly	
			65					70					75			

TOC	TGC	AAC	AAC	TAT	TANGCCCATG	CGGGCCCCAT	CCCGCGACCC	GGCATCGTCG	394
Ser	Cys	Asn	Asn	Tyr					
		90							

CCGGGG 400

			225
(i)	SEQUENCI	E CHARACTERISTI	CS:

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 151:

Met Arg Leu Ser Leu Thr Ala Leu Ser Ala Gly Val Gly Ala Val Ala

Met Ser Leu Thr Val Gly Ala Gly Val Ala Ser Ala Asp Pro Val Asp

Asn Ala Thr Asp Pro Gly Ala Ala Ala Gla Phe Asa Ala Ser Pro Val Als Glo Ser Tyr Leu Arg Asn Phe Lau Ala Ala Pro Pro Pro Glo Arg Ala Ala Met Ala Ala Gin Leu Gin Ala Val Pro Gly Ala Ala Gin Tyr

Ile Gly Leu Val Glu Ser Val Ala Gly Ser Cys Asn Asn Tyr

60

-5 Ala Val Ile Aso Tor Thr Cys Aso Tyr Gly Glo Val Val Ala Ala Leu

2.0

75

(2) INFORMATION FOR SEC TO NO: 152: (i) SECUENCE CHARACTERISTICS: (A) LENGTH: 990 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS; single (D) TOPOLOGY: linear (ii) BOLECULE TYPE: CONA

(B) TYPE: amino acid (D) TOPOLOGY; linear (ii) MOLECULE TYPE: protein

-28 -25

~3.0

55

70

(ix) FRATURE:	
(A) NAME/KEY: Coding Sequence	
(B) LOCATION: 93890	
(D) OTHER IMPORMATION:	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 152:	
AATAGTAATA TOOCTSTGCS GTTGCAAAAC GTSTSACCSA GGTTCCGCAG TCSAGCGCTG	60
CGGGCCGCCT TCGAGGAGGA CGAACCACAC TC ATG ACG AAC ATC GTG GTC CTG	113
Met Thr Asn Ile Val Val Len	
1 5	
ENT bad day day one are and and and one see one are day day	7.67

20

209

Ile Lys Gin Val Pro Asp Thr Trp Ser Glu Arg Lys Leu Thr Asp Gly 15

GAT TITC ACG CIG GAC COC GAS GCC GCC GAC GCG GTG CTG GAC GAS ATC

*	qes	Phe 25	The	Leu	Asp	Arg	Glu 30	Ala	Ala	Asp	Ala	V&1 35	Lsu	Asp	elu	He	
									CTA								257
*	46	Gáta	Arg	Ala	Vai	45	948	Aia	Leu	Gin	S0	Arg	Uau	riges	Gin	55 55	
									ACC								305
4	Ala	qsA	Gly	Ile	Glu 50	Gly	Ser	Val	Thr	Val 6S	Leu	Thr	Ala	Gly	Pro 70	Glu	
																AAG	353
,	Arg	Ala	Thr	75	Ala	Tie	Arg	Lys	86 28	Lec	Ser	Met	Gly	85 85	Asp	Lys	
									ATO								403
4	ALA	Val	His SO	Leu	Lys	Asp	QuA	95	Met.	His	Gly	Ser	100	Val	Tle	Gln	
									TTO								449
80	Thr	305	Trp	Ala	Leu	Ala	Arg 110	Ala	Leu	Gly	Thr	Tle	Glu	dly	Thr	Glu	
									ACC								497
	Leu 120	Val	Ile	Ala	Gly	Asn 125	Glu	Sex	Thr	Asp	130	Val	Giy	Gly		Val 135	
									GGC								545
9	ero.	Ala	Lie	Lia	140	Giu	zyx	Let)	Gly	145	STO	Gin	Leu	THE	150	Lett	
									AAG								593
á	yrg	Lys	Val	361 155	Iie	Glu	GTA	GIY	Lys 160	ile	Trix	GIĀ	Giu	165	G13	Thr	
									SCC								641
4	Amp	Gi i i	270		Phe	Tax	Leu	375	Ala	Tax.	Lea	Vxo	180	yaı	170	Ser	
									coc								689
	V8.1	185	310	nys	110	Asn	190	820	Arg	Phe	PEO	195	Pho	Lys	era	110	
									ACC								737
	290	Aia	ALA	uya	Lys	302 TAR	4117	vai	Thr	Agr	210	THY	nen	AIA	918	215	
									CTG								785
	GIY	Val	Slu	Sex	Asp 220	Glu	Val	gly	Leu	Ala 225	Asn	Ala	Gly	Ser	230	Val	
																ACC	833
	leu.	Ala	ser	235	PYO	Lys	ALO	Ala	Ly6 246	The	Ala	Cly	Gla	198 245	VAL	Thr	
																CAG	881
	qaa	Glu	Gly 250		Oly	Gly	Asn	03n 259		Val	Gln	Tyr	Leu 260	Val	Ala	Gln	

227 ARA ATC ATC TAAGACATAC GCACCTCCCA AAGACGAGAG CGATATAACC CATGGCTGA 939 Lvs Ile Ile 265 AGTACTOGTG CTCSTTGAGC ACCCTGAAGG CGCSTTAAAG AAGGTCAGCG C 990 (2) INFORMATION FOR SEC ID NO: 153: (1) SECURNCE CHARACTERISTICS: (A) LENGTH: 266 amino acids (8) TYPE: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (ii) MOLECULE TYPE: protein (v) FRAGMENT TYPE: internal (xi) SEQUENCE DESCRIPTION: SEG ID NO: 153: Met Thr Asm Ile Val Val Leu Ile Lys Glm Val Pro Asp Thr Trp Ser Giu Arg Lys Leu Thr Asp Gly Asp Phe Thr Leu Asp Arg Glu Ala Als Amp Ala Val Leu Amp Glu Tie Amp Glu Arg Ala Val Glu Glu Ala Leu 35 Gln Ile Arg Glu Lys Glu Ala Ala Asp Gly Ile Glu Gly Ser Val Thr Val Leu Thr Ala Gly Pro Glu Arg Ala Thr Glu Ala Ile Arg Lys Ala Leu Ser Met Gly Ala Asp Lys Ala Val His Leu Lys Asp Asp Gly Met His Gly Ser Asp Val Ile Gln Thr Gly Trp Ala Leu Ala Arg Ala Leu Gly Thr Ile Glu Gly Thr Glu Leu Val Ile Ala Gly Asn Glu Ser Thr 335 120 125 Asp Gly Val Gly Gly Ale Val Pro Ale Tie Ile Ale Glo Tyr Leu Gly 135 Leu Pro Glo Leu Thr Bis Lou Arg Lys Val Ser Tle Glu Gly Gly Lys Ile Thr Gly Glu Arg Glu Thr Asp Glu Gly Vol Phe Thr Lou Glu Ala 165 1.76 The Leu Pro Ala Val Ila Ser Val Ash Glu Lys Tie Ash Glu Pro Arg 185 Phe Pro Ser Phe Lys Gly Lie Met Ala Ala Lys Lys Lys Glu Val Thr 195 200 205

Val Leu Thr Leu Ala Shu lie Gly Val Glu Ser Asp Glu Val Gly Leu

20.0

									2	28						
	210					215					220					
Ala 225	Asm	Alo	gty	Ser	Thr 230	Val	Lesus	Ala	Ser	Thix 235	Pro	Lys	Pro	Ala	Lys 240	
Thr	Ala	Gly	Glu	Lye 245	Val	Thr	Asp	Glu	Gly 250	@lu	Gly	Gly	Asn	Gln 255	lle	
Val	Gln	Tyr	Leu 260	Val	Ala	Gln	Lys	Tle 265	Ile							
(2)	INFO	ORMA:	rion	FOR	SEQ	ID I	NO:	154:								
	(1)	\$2 \$3 \$5	i Li i T	CE CE ENGTE EPE: FRANC OPOLA	H: 2! DEDM	s ba leic ESS:	se pacio	sir» d								
	(xi	SE	OURS	CE D	ESCR.	IPTI	OW: :	SEQ	ID M	): 1.	94:					
CTG	GAT	TA:	rgaa	CCTA	CG G	CGCC										25
(2)	INF	DEMAC	TON	FOR	SEQ	ID :	SO:	155:								
	12	() ()	R) 11 8) T C) S	CS C ENGT: YPE: TRANS	B: 3' DWC DRDN	s ba l*ic RSS:	se p aci sin	airs d								
	1363	SB(	OUEN	CB D	SSCR	irti	ON:	SNQ :	io d	3: \$1	55.;					
CTC	CAT	GGT :	eccc.	TAGG.	AC C	0990	CAGO	c co	ggc							35
(2)	INF	ORMA	TION	FOR	\$8Q	ID:	100 t	156:								
	ξi	\$1 \$1	a) t c) s	CE C ENGT YPE: TRAN OPOL	H; 2 NUC DEDM	9 ba leic ESS:	se p aci sin	airs d								
	(sei	SP	QUEM	CS D	escr	ipti	ON:	880	ID N	0: 1:	96:					
CRE	MAT	CTA '	TGAG	ectg	TC G	TTGA	ccac									29
(2)	INY	ÖRMA	TION	POR	SEQ	ID	NO:	157:								
	(i	0	A) L B) T C) S	CE C ENGT YPL: TRAN OPOL	H: 3 nuc erdn	0 ba leic ESS:	se p aci sin	airs d								

(xi) SEQUENCE DESCRIPTION: SNQ ID NO: 157:	
CTCCCCGGGC TTANTAGTTG TTGCRGGAGC	30
(2) INFORMATION FOR SEQ ID NO: 158:	
(i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 32 hase pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear	
(*i) REQUENCE DESCRIPTION: SEQ ID NO: 158:	
GCTTAGATCT ATGATTTTCT GGGCAACUAG GTA	33
(2) INFORMATION FOR SEQ ID NO: 159:	
(i) SEQUENCE CHARACTERISTICS: (A) LEMOTH: 30 base pairs (B) TTPE: muclaic acid (C) STRANDEDNESS: single (D) TOFOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 159:	
GCTTCCATGG GCGAGCCACA GGCGTGGGRA	30
(2) IMPORMATION FOR SEQ ID NO: 160:	
(i) SECHENCE CHARACTERISTICS: (A) LENGTH: 30 base paixs (B) TFFE: nucleic acid (C) STRANDERNESS: single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 160:	
CYGAGATCTA GAATGCCACA GGGAACTGTG	30
(2) IMPORMATION FOR SEQ ID NO: 161:	
(i) SEQUENCE CHARACTERISTICS: (A) LEMOTH: 30 base pairs (B) TYPE: mucleic acid (C) STRANDEDNESS: Single (D) TOPOLOGY: linear	
(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 161:	
TOTOCCGGGG GTAACTUAGA GCGAGCGGAC	36
(2) INFORMATION FOR SEC IS NO: 162:	

	(i) SEQUENCS CHARACTERISTICS: (A) LENGTH; 27 base pairs (B) TYPE; THOUGH acid (C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 162;	
CTGI	AGATOTA TGAACGTCAC CGTATCC	21
{2}	IMPORMATION FOR SEQ ID NO: 163:	
	(i) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 27 base pairs	
	(B) TYPE: nucleic acid	
	(C) STRANDEDMESS: single	
	(D) TOPOLOGY: linear	
	(mi) SEQUENCE DESCRIPTION: SEQ ID NO: 163:	
rgs/repy	CCCGGGG CTCACCCACC GGCGACG	25
2021	ceron crace occare	
(2)	INFORMATION FOR SEQ ID NO: 164:	
	(1) SEQUENCE CHARACTERISTICS:	
	(A) LENGTH: 30 base pairs	
	(B) TYFE: nucleic acid	
	(C) STRANDERNESS: single	
	(D) TOPOLOGY: linear	
	(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 164:	
CARR	AGATCTA TGGCAACACG TTTTATGACG	.30
(2)	IMPORMATION FOR SEQ ID NO: 165:	
	(1) SEQUENCE CHARACTERISTICS:	
	(A) LEMGTH: 30 base pairs	
	(B) TYPE: numletc soid	
	(C) STRANDEDNESS: single	
	(D) TOPOLOGY: linear	
	(mi) SEQUENCE DESCRIPTION: SEQ ID NO: 165:	
CTC	CCCODET TAGCTECTOA OBATCTECTN	36
(2)	IMPORMATION FOR SEQ ID NO: 166:	
	(x) SEQUENCE CHARACTERISTICE:	
	(A) LEMOTH: 31 base pairs	
	(%) TYPE: nucleic acid	
	(C) STRANDEDNESS: single	
	(D) TOPOLOGY limear	

231

(wi) SECURACE DESCRIPTION: SEO ID NO: 166: CITGARGATCT ATGCCCAAGA GAAGCGAATA C 31 (2) INFORMATION FOR SEC ID NO: 167: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 31 base pairs (B) TYPE: nucleic acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 167: CGGCAGCTGC TAGCATTCTC CGAATCTGCC G 31 (2) INFORMATION FOR SEQ ID NO: 168: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 15 amino acids (B) TYPE: amino acid (C) STRANDEDNESS; single (D) TOPOLOGY: linear (11) MOLECULE TYPE: None (xi) SEQUENCE DESCRIPTION: SEO ID NO: 168: Pro Gin Gly Thr Val Lys Trp Phe Asn Ala Glu Lys Gly Phe Gly 3 3.0 (2) IMPORMATION FOR SEQ ID NO: 169: (i) SEQUENCE CHARACTERISTICS: (A) LENGTH: 15 amino acids (3) TYPS: amino acid (C) STRANDEDNESS: single (D) TOPOLOGY: linear (11) MOLECULE TYPE: None (ix) FEATURE: (A) NAME/KEY: Other (B) LOCATION: 15 (D) OTHER INFORMATION: Raa is unknown (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 169: Asn Val Thr Val Ser Ile Pro Thr Ile Leu Arg Pro Xaa Xaa Xaa 10 (2) INFORMATION FOR MEQ ID NO: 170:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 15 amino acids

(B) TYPE: amino acid

(C) STRANDEDNESS: single

232

- (D) TOPOLOGY: linear
- (ii) MOLECULE TYPE: None
- (ix) FEATURE:
  - (Al NAME/KEY: Other
  - (B) LOCATION: 1
  - (D) OTHER INFORMATION: The Could also be Ala
- (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 176:

Thr Arg Phe Met Thr Asp Pro His Ala Met Arg Asp Met Ala Gly 1 5 10 15

- (2) INFORMATION FOR SEQ ID NO: 171:
  - (i) SEQUENCE CHARACTERISTICS:
    - (A) LENGTH: 15 amino acids
      - (8) TYPE: amino acid
      - (C) STRANDEDNESS: single
      - (D) TOPOLOGY: linear
  - (ii) MOLECULE TYPE: None
  - (xi) SEQUENCE DESCRIPTION: SEQ TO NO: 171:

Pro Lys Arg Ser Glu Tyr Arg Gln Gly The Pro Aso Trp Val Asp 1 5 10 15

- (2) INFORMATION FOR SEQ ID NO:172:
  - (1) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 404 amino acids
    - (B) TYPE: amino acid
    - (C) STRANDEDNESS: single
    - (D) TOPOLOGY: linear
  - (xi) SEQUENCE DESCRIPTION: SEQ 10 NO:172:

Mot Ala Tor Val Asn Arg Ser Arg His His His His His His His His 1 10 Ile Glu Gly Arg Ser Phe Ser Arg Pro Gly Lew Pro Val Glu Tyr Lew 25 Gln Val Pro Ser Pro Ser Met Gly Arg Asp 11s Lys Val Cln Phe Gln 45 Ser Gly Gly Asn Asn Ser Pro Ala Val Tyr Leu Leu Asp Gly Leu Arg Ala Gin Asp Asp Tyr Asn Gly Txp Asp Ile Asn Thr Pro Ala Phe Glu Trp Tyr Tyr Gln Sex Gly Leu Ser Ile Val Met Pro Val Gly Gly Gln 98 Ser Ser Phe Tyr Ser Asp Trp Tyr Ser Pro Ala Cye Gly Lys Ala Gly 1.05 110 Cys Sic Thr Tyr Lys Trp Slu Thr Pne Leu Thr Ser Glu Leu Pro Sin 128

Trp Leu Ser Als Asu Ard Ala Val Lvs Pro Thr Clv Ser Ala Ala Ile

233

1.35 Gly Leu Ser Met Ala Gly Ser Ser Ala Met Ile Leu Ala Ala Tyr His 150 155 Pro Gin Gin Phe Ile Tyr Als Gly Ser Lou Ser Ala Leu Leu Asp Pro 165 176 Ser Gln Gly Met Gly Pro Ser Leu Tie Gly Leu Ala Met Gly Asp Ala 185 Gly Gly Tyr Lys Ala Ala Asp Met Trp Gly Pro Ser Ser Asp Pro Ala 195 200 Trp Glu Arg Asn Asp Pro Thr Gln Gln Ile Pro Lys Leu Val Ala Asn 215 Asn Thr Arg Leu Trp Val Tyr Cys Gly Asn Gly Thr Pro Asn Glu Leu 230 238 Gly Gly Ala Asn Ile Pro Ala Glu Phe Leu Glu Asn Phe Val Arg Ser 245 250 Ser Asn Leu Lys Phe Gln Asp Ala Tyr Asn Ala Ala Gly Gly His Asn 265 Ala Val Phe Asn Phe Pro Pro Asn Gly Thr His Ser Trp Glu Tyr Trp 286 285 Gly Ala Gin Leu Asa Ala Met Lys Gly Asp Leu Gin Ser Ser Leu Gly 295 380 Ala Gly Lys Leu Ala Met Thr Glu Gln Gln Trp Asn Phe Ala Gly Ile 320 315 Glu Ala Ala Ala Ser Ala Ile Gin Gly Asn Val Thr Ser Ile His Ser 325 330 Leu Leu Asp Clu Gly Lys Gln Ser Leu Thr Lys Leu Ala Ala Ala Trp 340 345 Gly Gly Ser Gly Ser Glu Ala Tyr Gln Gly Val Gln Gln Lys Trp Asp 360 Ala Thr Ala Thr Glu Leo Asn Asn Ala Leu Gln Asn Leu Ala Arg Thr 378 3.80 Ile Ser Glu Ala Gly Gin Ala Met Ala Ser Thr Glu Gly Asn Val Thr 3.95 Gly Met Phe Ala

## (2) INFORMATION FOR SEQ ID NO:173:

- (i) SEQUENCE CHARACTERISTICS:
  - (A) LENGTH: 403 amino acids
  - (B) TYPE: amino soid
  - (C) STRANDEDNESS: single
  - (D) TOPOLOGY: linear

# (xi) SEQUENCE DESCRIPTION: SEQ ID NO:173:

				85					90					95		
110	Sex	Glu		Gly	Gln	Ala	Mot			Thr	Gits	Gly	Asa	Van 2	Thr	
			300					105					110			
Gly	Met	phe	RIA	Lyg	Leu	Phe	Ser	YL,G	850	Gly	1.0.0	Pro	V&l	Glu	Tyr	
		115					120					328				
Leu	Gla	Val	\$20	Ser	Pro	Ser	Met	dly	Arg	Asp	lle	Lya	Val	Gin	Phe	
	130					135					140					
Gla	Sex	GLY	Gly	Asn	Asn	Set	Pro	Ala	Va.	Tyr	£18373	SAFE	Asp	Gly	Less	
145					150					155					1.60	
Arg	Ala	Gln	RHO	Asp	Tyr	Ass	Gly	Trp	Asp	Hie	Asn	Thr	Pro	Ala	Phe	
				165					170					275		
Glu	Tro	Tyr	Tyr	Gin	Ser	Gly	Let	Ser	Tle	Val.	Met	Pro	Val	Gly	Gly	
			180					185					190			
Gla	Ser	Ser	Phe	Tyr	ser	Asp	Trp	Tyr	Ser	Pro	Ala	Cys	Gly	Lys	Ala	
		195					200					205				
Gly	Cys	Glm	Thr	Tyr	lays	Trp	Glu	The	Phie	Leu	Thr	Ser	Glu	Leu	Pro	
	210					215					220					
Gla	Trp	Leu	Ser	Ala	Asn	Arg	Ala	Val	Lys	Pro	Thr	Gly	Ser	Ala	Ala	
225					230					235					240	
Ile	Gly	Lens	Ser	Met	Ala	Gly	Ser	Sex	Ala	Met	110	Lett	Ala	Ala	Tyr	
				245					250					255		
His	Pro	Glm	Glm	Phe	110	Tyx	Ala	Gly	Ser	Leu	Sex	Ala	Leu	Leu	Rap	
			260					265					275			
Pro	Sex	Glin	Gly	Men	Gly	Pro	Sex	1.00	Lle	Gly	Leu	Ala	Me t	Gly	Asp	
		275					280					285				
Ala	Gly	Gly	Tyr	Ze s	Ala	Ala	Asp	36e t	Tro	GLy	Pro	805	Ser	Augs	Pro	
	290					298					300					
Ala	Trp	Glu	Arg	Asn	Asp	Pro	Thr	Gla	Gln	Tie	Pro	Lys	Leu	Val	Ala	
305					310					315					320	
Asn.	Asn	Thr	Arg	Leo	Trp	Val	Tyr	Cys	Gly	asn	dly	2752	220	Asn	Glu	
				325					330					335		
Leu	Gly	Gly	Ala	Asn	lle	2ro	Ala	Glu	Phe	Leu	Glu	Aso	2430	Val	Arg	
			340					345					350			
Ser	Ser	Assi	Letta	Lys	Piso	Gla	Assp	Ala	Tyr	Asses	Ala	Ala	Gly	Gly	13 i. st	
		355					360					365				
Asn	Ala	VAL	Phe	Asn	Phe	320	Pro	Agn	Gly	Thr	Bis	362	Try	Glu	Tyr	
	370					375					380					
TYP	Gly	Ala	Gin	Leu	Asn	Ala	Meto	Lys	Gly	Asp	2,650	Gln	Ser	ser	Leu	
385					390					395					400	
Glv	Ala	Gly														

WO 98/44119

235

CLAIMS

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1. A substantially pure polypeptide fragment which

a) comprises an amino acid sequence selected from the sequences shown in SEQ ID NO: 2, 4, 6, 8, 10, 12, 14, 16, 17-23, 42, 48, 50, 52, 54, 56, 58, 60, 62, 64, 66, 68, 70, 72-86, 88, 90, 92, 94, 141, 143, 145, 147, 149, 151, 153, and 168-171,

PCT/DK98/00132

- b) comprises a subsequence of the polypeptide fragment
  defined in a) which has a length of at least 6 amino
  acid residues, said subsequence being immunologically
  equivalent to the polypeptide defined in a) with
  respect to the ability of evoking a protective immune
  response against infections with mycobacteria belonging
  to the tuberculosis complex or with respect to the
  ability of eliciting a diagnostically significant
  immune response indicating previous or ongoing sensitization with antigens derived from sycobacteria belonging to the tuberculosis complex, or
- e3 comprises an amino acid sequence having a sequence 20 identity with the polypeptide defined in a) or the subsequence defined in b) of at least 70% and at the same time being immunologically equivalent to the polypeptide defined in a) with respect to the ability of evoking a protective immune response against infec-25 tions with mycobacteria belonging to the tuberculosis complex or with respect to the ability of eliciting a diagnostically significant immune response indicating previous or ongoing sensitization with antigens derived from mycobacteria belonging to the tuberculosis com-30 plex,

with the proviso that

i) the polypeptide fragment is in essentially pure form when consisting of the amino acid sequence 1-96 of SEQ ID NO: 2 or when consisting of the amino acid sequence 87-108 of SEQ ID NO: 4 fused to \$0-galactosidase,

- 5 ii) the degree of sequence identity in c) is at least 95% when the polypeptide comprises a homologue of a polypeptide which has the amino acid sequence SEQ ID NO: 12 or a subsequence thereof as defined in b), and
- iii) the polypeptide fragment contains a threonine residue corresponding to position 213 in SEC ID NO: 42 when comprising an amino acid sequence of at least 6 amino acids in SEO ID NO: 42.
  - 2. The polypeptide fragment according to claim 1 in essentially pure form.
- 3. The polypeptide fragment according to claim 1 or 2, which comprises an epitope for a T-helper cell.
  - 4. The polypeptide fragment according to any of the preceding claims, which has a length of at least 7 amino acid residues, such as at least 8, at least 9, at least 10, at least 12, at
- 20 least 14, at least 16, at least 18, at least 20, at least 22, at least 24, and at least 30 amino acid residues.
  - The polypeptide framment according to any of the preceding claims, which is free from amino acid residues -30 to -1 in SEQ ID NO: 6 and/or -32 to -1 in SEQ ID NO: 10 and/or -8 to
- 25 -1 in SEO ID NO: 12 and/or -32 to -1 in SEO ID NO: 14 and/or -33 to -1 in SEO ID NO: 42 and/or -38 to -1 in SEO ID NO: 52 and/or -33 to -1 in SEO ID NO: 56 and/or -56 to -1 in SEO ID NO: 58 and/or -28 to -1 in SEO ID NO: 151.
- 6. The polypeptide fragment according to any of the preceding 36 claims which is free from any signal sequence.

7. The polypeptide fragment according to any of the preceding claims which

- 1) induces a release of IFN-\(\gamma\) from primed memory T-lymphocytes withdrawn from a mouse within 2 weeks of primary infection or within 4 days after the mouse has been rechallenge infected with mycobacteria belonging to the tuberculosis complex, the induction performed by the addition of the polypeptide to a suspension comprising about 200.000 spleen cells per ml, the addition of the polypeptide resulting in a concentration of 1-4 \(\mu\)g polypeptide per ml suspension, the release of IFN-\(\gamma\) being assessable by determination of IFN-\(\gamma\) in supernatant harvested 2 days after the addition of the polypeptide to the suspension, and/or
- 15 2) induces a release of IFN-y of at least 300 pg above background level from about 1000,000 human PBMC (peripheral blood mononuclear cells) per ml isolated from TB patients in the first phase of infection, or from healthy BCG vaccinated donors, or from healthy contacts to TB patients, the induction being performed by the 20 addition of the polypeotide to a suspension comprising the about 1,000,000 PEMC per ml, the addition of the polypeptide resulting in a concentration of 1-4 µg polypeptide per ml suspension, the release of IFN-v 25 being assessable by determination of IFN-y in supermatant harvested 2 days after the addition of the polypeptide to the suspension; and/or
- 3) induces an IPN-y release from bovine PBMC derived from animals previously sensitized with mycobacteria belonging to the tuberculosis complex, said release being at least two times the release observed from bovine PBMC derived from animals not previously sensitized with mycobacteria belonging to the tuberculosis complex.

238

- 8. A polypeptide fragment according to any of the preceding claims, wherein the sequence identity in c) is at least 80%, such as at least 85%, at least 90%, at least 91%, at least 92%, at least 93%, at least 94%, at least 95%, at least 96%, at least 93%, at least 93%, at least 99%, and at least 95.5%.
  - A fusion polypeptide comprising at least one polypeptide fragment according to any of the preceding claims and at least one fusion partner.
- 10. A fusion polypeptide according to claim 56, wherein the fusion partner is selected from the group consisting of a polypeptide fragment as defined in any of claims 1-8, and an other polypeptide fragment derived from a bacterium belonging to the tuberculosis complex, such as ESAT-6 or at least one T-cell epitope thereof, MPB64 or at least one T-cell epitope thereof, MPT64 or at least one T-cell epitope thereof, and MPB59 or at least one T-cell epitope thereof.

# 11. A fusion polypeptide fragment which comprises

- 1) a first amino acid sequence including at least one stretch of amino acids constituting a T-cell epitope derived from the M. tuberculosis protein ESAT-6, and a second amino acid sequence including at least one T-cell epitope derived from a M. tuberculosis protein different from ESAT-6 and/or including a stretch of amino acids which protects the first amino acid sequence from in vivo degradation or post-translational processing; or
- 2) a first amino acid sequence including at least one stretch of amino acids constituting a T-cell epitope derived from the M. tuberculosis protein MPT59, and a second amino acid sequence including at least one T-cell epitope derived from a M. tuberculosis protein different from MPT59 and/or including a stretch of amino acids which protects the first amino acid

> sequence from in vivo degradation or post-translational processing.

- 12. A fusion polypeptide fragment according to claim 11, wherein the first amino acid sequence is situated C-termi-5 nally to the second amino acid sequence.
  - 13. A fusion polypeptide fragment according to claim 11, wherein the first amino acid secuence is situated N-terminally to the second amino acid sequence.
- 14. A fusion polypeptide fragment according to any of claims 10 11-13, wherein the at least one T-cell epitope included in the second amino acid sequence is derived from a M. tuberculosis polypeptide selected from the group consisting of a polypeptide fragment according to any of claims 1-55, Dnak, GroEL, urease, glutamine synthetase, the proline rich com-
- 15 plex, L-alanine dehydrogenase, phosphate binding protein, Ag 85 complex, HBHA (heparin binding hemagglutinin), MFT51, MPT64, superoxide dismutase, 19 kDa lipoprotein, o-crystallin, Grows, MPT59 when the first T-cell epitope is derived from BSAT-6, and BSAT-6 when the first T-cell epitope is
- 20 derived from MPTS9.
  - 15. A fusion polypeptide fragment according to any of claims 11-14, wherein the first and second T-cell epitopes each have a sequence identity of at least 70% with the natively occurring sequence in the proteins from which they are derived.
- 25 16. A fusion polypeptide according to any of claims 11-15, wherein the first and/or second amino acid sequence have a sequence identity of at least 70% with the protein from which they are derived.
- 17. A fusion polypeptide fragment according to any of claims 30 11-16, wherein the first amino acid sequence is the amino acid secuence of ESAT-6 or of MPT59 and/or the second amino acid sequence is the amino acid sequence of a M. tuberculosis

that of BSAT-6.

WO 98/44119 PCT/DK98/00132

polypeptide selected from the group consisting of a polypeptide fragment according to any of claims 1-8, DnaK, GroEL, urease, glutamine synthetase, the proline rich complex, L-alanine dehydrogenase, phosphate binding protein, Ag 5 85 complex, BBHA (heparin binding hemagglutinin), MPT51, MPT64, superoxide dismutase, 19 kDa lipoprotein, α-crystallin, GroES, ESAT-6 when the first amino acid sequence is that of MPT59, and MPT59 when the first amino acid sequence is

- 10 18. A fusion polypeptide fragment according to any of claims 11-17, which comprises ESAT-6 fused to MPT59.
  - 19. A fusion polypeptide fragment according to claim 18, wherein no linkers are introduced between the two amino acid sequences.
- 15 20. A polypeptide according to any of the preceding claims which is lipidated so as to allow a self-adjuvating effect of the polypeptide.
  - 21. A substantially pure polypeptide according to any of claims 1-20 for use as a pharmaceutical.
  - 20 22. The use of a substantially pure polypeptide according to any of claims 1-20 in the preparation of a pharmaceutical composition for the diagnosis of or vaccination against tuberculosis caused by Mycobacterium tuberculosis, Mycobacterium africanum or Mycobacterium bovis.
  - 25 23. A nucleic acid fragment in isolated form which
    - comprises a nucleic acid sequence which encodes a polypeptide as defined in any of claims 1-20, or comprises a nucleic acid sequence complementary thereto.
  - has a length of at least 10 nucleotides and hybridizes
     readily under stringent hybridization conditions with a

243

nucleic acid fragment which has a nucleotide sequence selected from SEQ ID NO: 1 or a sequence complementary thereto, SEQ ID NO: 3 or a sequence complementary thereto, S SEQ ID NO: 5 or a sequence complementary thereto. SEQ ID NO: 7 or a sequence complementary thereto, SEQ ID NO: 9 or a sequence complementary thereto, SEQ ID NO: 11 or a sequence complementary thereto, SEO ID NO: 13 or a sequence complementary thereto, SEQ ID NO: 15 or a sequence complementary thereto, 10 SEQ ID NO: 41 or a sequence complementary thereto, SEC ID NO: 47 or a sequence complementary thereto, SEC ID No: 49 or a sequence complementary thereto, SEC ID NO: 51 or a sequence complementary thereto, 15 SEC ID NO: 53 or a sequence complementary thereto, SEC ID NO: 55 or a sequence complementary thereto, SEQ ID NO: 57 or a sequence complementary thereto, SEQ ID NO: 59 or a sequence complementary thereto, SEC ID NC: 61 or a sequence complementary thereto, 20 SEQ ID NO: 63 or a sequence complementary thereto, SEQ ID NO: 65 or a sequence complementary thereto, SEO ID NO: 67 or a sequence complementary thereto, SEC ID NO: 69 or a sequence complementary thereto, SEO ID NO: 71 or a sequence complementary thereto, 25 SEG ID NO: 87 or a sequence complementary thereto, SEO ID NO: 89 or a sequence complementary thereto, SEQ ID NO: 91 or a sequence complementary thereto, SEO ID NO: 93 or a sequence complementary thereto, SEC ID NO: 140 or a sequence complementary thereto, 30 SEQ ID NO: 142 or a sequence complementary thereto, SEC ID NO: 14% or a sequence complementary thereto, SEQ ID NO: 146 or a sequence complementary thereto, SEO TD NO: 148 or a secuence complementary thereto. SEQ ID NO: 150 or a sequence complementary thereto, and 35 SEO ID NO: 152 or a sequence complementary thereto.

with the provise that when the nucleic acid fragment comprises a subsequence of SEQ ID NO: 41, then the nucleic acid

fragment contains an A corresponding to position 781 in SEQ ID NO: 41 and when the nucleic acid fragment comprises a subsequence of a nucleotide sequence exactly complementary to SEQ ID NO: 41, then the nucleic acid fragment comprises a T corresponding to position 781 in SEQ ID NO: 41.

242

- 24. A nucleic acid fragment according to claim 23, which is a DNA fragment.
- 25. A vaccine comprising a nucleic acid fragment according to claim 23 or 24, the vaccine effecting in vivo expression of antigen by an animal, including a human being, to whom the vaccine has been administered, the amount of expressed antigen being effective to confer substantially increased resistance to infections with mycobacteria of the tuberculosis complex in an animal, including a human being.
- 15 26. A nucleic acid fragment according to claim 23 or 24 for use as a pharmaceutical.
  - 27. The use of a nucleic acid fragment according to claim 23 or 24 in the preparation of a pharmaceutical composition for the diagnosis of or vaccination against tuberculosis caused by Mycobacterium tuberculosis, Mycobacterium africanum or Mycobacterium bovis.
    - 28. An immunologic composition comparising a polypeptide according to any of claims 1-20.
  - 29. An immunologic composition according to claim 28, which further comprises an immunologically and pharmaceutically acceptable carrier, vehicle or adjuvant.
    - 30. An immunologic composition according to claim 29, wherein the carrier is selected from the group consisting of a polymer to which the polypeptide(s) is/are bound by hydrophobic
  - 30 non-covalent interaction, such as a plastic, e.g. polystyrene, a polymer to which the polypeptide(s) is/are covalently

bound, such as a polysaccharide, and a polypeptide, e.g. bovine serum albumin, ovalbumin or keyhole limpet hemocyanin; the vehicle is selected from the group consisting of a diluent and a suspending agent; and the adjuvant is selected from 5 the group consisting of dimethyldioctadecylammonium bromide (DDA), Quil A, poly I:C, Freund's incomplete adjuvant, IFN-γ, IL-2, IL-12, monophosphoryl lipid A (MPL), and muramyl dipeptide (MDP).

- 31. An immunologic composition according to any of claims 28 10 to 30, comprising at least two different polypeptide fragments, each different polypeptide fragment being a polypeptide according to any of claims 1-20.
- 32. An immunologic composition according to claim 31, comprising 3-20 different polypeptide fragments, each different polypeptide fragment being according to any of claims 1-20.
  - 33. An immunologic composition according to any of claims 28-32, which is in the form of a vaccine.
    - 34. An immunologic composition according to any of claims 28-
    - 32, which is in the form of a skin test reagent.
- 20 35. A vaccine for immunizing an animal, including a human being, against tuberculosis caused by mycobacteria belonging to the tuberculosis complex, comprising as the effective component a non-pathogenic microorganism, wherein at least one copy of a DNA fragment comprising a DNA sequence encoding a polypeptide according to any of claims 1-20 has been incorporated into the genome of the microorganism in a manner allowing the microorganism to express and optionally secrete the polypeptide.
- 36. A vaccine according to claim 35, wherein the microorga-30 nism is a bacterium.

244

37. A vaccine according to claim 36, wherein the bacterium is selected from the group consisting of the genera Mycobacterium. Salmonella. Fseudomonas and Eschericia.

- 38. A vaccine according to claim 37, wherein the microorga-5 nism is Mycobacterium bovis BCG, such as Mycobacterium bovis BCG strain: Danish 1331.
- 39. A vaccine according to any of claims 35-38, wherein at least 2 copies of a DNA fragment encoding a polypeptide according to any of claims 1-20 are incorporated into the genome of the microorganism.
  - 40. A vaccine according to claim 39, wherein the number of copies is at least 5.
  - 41. A replicable expression vector which comprises a nucleic acid fragment according to claim 23 or 24.
- 15 42. A vector according to claim 41, which is selected from the group consisting of a virus, a hacteriophage, a plasmid, a cosmid, and a microchromosome.
  - 43. A transformed cell harbouring at least one vector according to claim 41 or 42.
- 20 44. A transformed cell according to claim 43, which is a bacterium belonging to the tuberculosis complex, such as a M. tuberculosis bovis BCG cell.
  - 45. A transformed cell according to claim 43 or 44, which expresses a polypeptide according to any of claims 1-20.
- 25 46. A method for producing a polypeptide according to any of claims 1-20, comprising

inserting a nucleic acid fragment according to claim 23 or 24 into a vector which is able to replicate in a host cell,

245

introducing the resulting recombinant vector into the host cell, culturing the host cell in a culture medium under conditions sufficient to effect expression of the polypeptide, and recovering the polypeptide from the host cell or culture medium; or

isolating the polypeptide from a short-term culture filtrate as defined in claim 1: or

isolating the polypeptide from whole mycobacteria of the tuberculosis complex or from lysates or fractions thereof.

10 e.g. cell wall containing fractions; or

synthesizing the polypeptide by solid or liquid phase peptide synthesis.

- 47. A method for producing an immunologic composition according to any of claims 28-32 comprising
- preparing, synthesizing or isolating a polypeptide according to any of claims 1-20, and

solubilizing or dispersing the polypeptide in a medium for a vaccine, and

optionally adding other M. tuberculosis antigens and/or a carrier, vehicle and/or adjuvant substance,

OX.

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cultivating a cell according to any of claims 37-45, and

transferring the cells to a medium for a vaccine, and

25 optionally adding a carrier, vehicle and/or adjuvant substance.

- 48. A method of diagnosing tuberculosis caused by Mycobacterium tuberculosis, Mycobacterium africanum or Mycobacterium bovis in an animal, including a human being, comprising intradermally injecting, in the animal, a polypeptide according to any of claims 1-20 or an immunologic composition according to claim 34, a positive skin response at the location of injection being indicative of the animal having tuberculosis, and a negative skin response at the location of injection being indicative of the animal not having tuberculosis.
- 49. A method for immunising an animal, including a human being, against tuberculosis caused by mycobacteria belonging to the tuberculosis complex, comprising administering to the animal the polypeptide according to any of claims 1-20, the immunologic composition according to claim 33, or the vaccine according to any of claims 35-40.
  - 50. A method according to claim 49, wherein the polypeptide, immunologic composition, or vaccine is administered by the parenteral (such as intravenous and intraasterially), intraperitoneal, intramuscular, subcutaneous, intradermal, oral, buccal, sublingual, nasal, rectal or transdermal route.

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- 51. A method for diagnosing ongoing or previous sensitization in an animal or a human being with bacteria belonging to the tuberculosis complex, the method comprising providing a blood sample from the animal or human being, and contacting the sample from the animal with the polypeptide according to any of claims 1-20, a significant release into the extracellular phase of at least one cytokine by mononuclear cells in the blood sample being indicative of the animal being sensitized.
- 30 52. A composition for diagnosing tuberculosis in an animal, including a human being, comprising a polypeptide according to any of claims 1-20, or a nucleic acid fragment according to claim 23 or 24, optionally in combination with a means for desection.

247

53. A monoclonal or polyclonal antibody, which is specifically reacting with a polypeptide according to any of claims 1-20 in an immuno assay, or a specific binding fragment of said antibody.

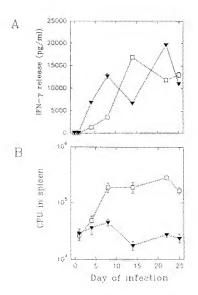


Fig. 1
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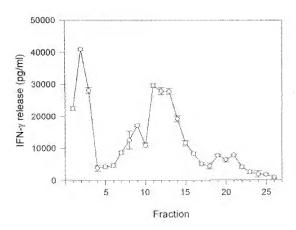


Fig. 2

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1	GAATTCGCCGGGTGCACACAGOCTTACACGACGC <u>AGC</u> TGGACACATGAAG M R	56
53	GGTCGGTCGGCGCTGCTGCGGGCGCTCTGGATTGCCGCACTGTCATTCGG	400
0.4	G R S A L L R A L W I A A L S F G	100
101	GTTGGGCGTGTCGCGGTAGCCGCGGGAACCCACCGCCAAGGCCGCCCCAT	1.50
	LGGVAVAAEPTAKAAP	200
151	ACGAGAACCTGATGGTGCCGTCGCCCTCGATGGGCCGGGACATCCCGGTG	200
	Y E N L M V P S P S M G R D I P V	200
201	GCCTTCCTAGCCGGTGGGCCGCACGCGGTGTATCTGCTGGACGCCTTCAA	250
	AFLAGGPHAVYLLDAFN	238
251	CGCCGGCCCGGATGTCAGTAACTGGGTCACCGCGGGTAACGCGATGAACA	ngn
20.52		300
201		0.00
301	CGTPGGCGGCAAGGGGATTTCGGTGGTGGCACCGGCCGGTGGTGCGTAC	350
Nn.	TLAGKGISVVAPA*GGAY	
303	AGCATGTACACCAACTGGGAGCAGGGAGCAGCAGCAGCAGCAGCAGCAGCAGCAGC	490
	S M Y T N W E Q D G S K Q W D T F	
401	CTTGTCCGCTGAGCTGCCCGACTGGCCGCTAACCGGGGCTTGGCCC	450
	LSAELPDWLAANRGLA	
451	CCGGTGGCCATGCGGCCGTTGGCGCCGCTCAGGGGGGGGTTACGGGGCGATG	560
	PGGHAAVGAAQGGYGAM	
501	GCGCTGGCGGCCTTCCACCCCGACCGCTTCGCCTCGCTCG	556
	A L A A F H P D R F G F A G S M S	
551	GGGCTTTTTGTACCCGTCGAACACCACCACCACCGGGGGGGG	500
	G F L Y P S N T T T N G A I A A	
601	GCATGCAGCAATTCGGCGGTGTGGACACCAACGGAATGTGGGGGAGCACCA	6.50
	G M Q Q F G G V D T N G M W G A P	
651	CAGCTGGGTCGGTGGAAGTGGCACGACCCGTGGGTGCATGCCAGCCTGCT	700
	Q I G R W R W B D P W V H A S L I	
701	GGCGCAAAACACCCGGGTGTGGGGTGTGGAGCCCGACCAACCCGGGAG	750
	A Q N N T R V W V W S P T N P G	
751	CCAGCGATCCCGCCGCCATGATCGGCCAAACCGCCGAGGCGATGGGTAAC	800
	ASDPAAMIGQIAEAMGN	00.
801	AGCCGCATGTTCTACAACCAGTATCGCAGCGTCGGCGGCAGAACGGACA	850
V-9-X	S R M F Y N Q Y R S V G G H N G H	2000
851	CTTCGACTTCCCAGCCAGCGGTGACAACGGCTGGGGGCTCGTGGGCGCCCC	900
	F D F P A S G D N G W G S W A P	Carles .
961	AGCTGGGCGCTATGTCGGGCGATATCGTCGGTGCGATCCGCTAAGCGAAT	956
4000	Q I G A M S G D I V G A I R	2.30
951		992
201	3 Au	3 76

Fig. 5

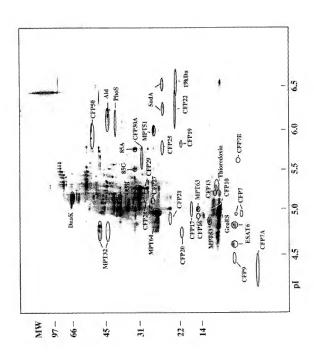


Fig. 6
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	see the claims see abstract; examples 1,3 see page 12 - page 18 see page 21 - page 25	nfnn		
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Box	Observations where certain claims were found unsearchable (Continuation of item 1 of first sheet)
This into	emational Search Report has not been established in respect of certain claims under Article 17(2)(4) for the following reasons:
1. X	Claims Nos : Decembe they relate to subject matter not required to be searched by this Authority, namely:
	Although claims 49 and 50 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.
s 🗀	Claims No.:  Necessary Buy visits in parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful international Search can be carried out, specifically.
3. [	Claims Nos." Secause www ure dependent claims and are not drahed in accordance with the second and third sentence of Fule 6.4(a).
Box II	Observations where unity of invention is tacking (Continuation of item 2 of first sheet)
Tries inte	emational Searching Authority found multiple inventions in this international application, as follows:
	see additional sheet
3. []	As all required additional search fees were briefly paid by the applicant, this informational Search Report covers at searchable claims.
2. [_]	As all seamonable claims sticid be searched without other pastering an additional less, this Authority sid not revise payment of any additional less.
3. X	As anly asome of the required additional search free were timely paid by the applicant, this international Search Report occurs only those claims for which free were part, specifically claims for.
	1-4,6-17,20-53; inventions 1 and 8
4 🗆	No reculted additional search fees were treely pricitly; the applicant. Consequently, this informational Search Report is restricted to the invention first mentioned in the oligane, it is covered by oberes Nov.
Homark	on Protest The additional search fees were accompanied by the applicant's protest.
	X) No protest accompanied the payment of additional yearth fees

### FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-4. 6-17. 20-53 all partially

A polypeptide fragment from mycobacteria belonging to the tuberculosis complex comprising the amino acid SEQ ID NO: 2, nucleic acids endoding said polypeptide as in SEQ ID NO: 1, fusion proteins comprising said polypeptides, vaccines, pharmaceutical and immunological compositions containing said polypeptide or nucleic acid, an expression vector comprising said nucleic acid, a host transformed with said vector, immunization with said polypeptide, the use of said polypeptide in diagnosis, antibodies against said polypeptide.

- Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEO ID NO: 4 and 3.
- Claims: 1-17, 20-53 all partially
   same as invention 1 but for SEO 1D NO: 6, 5 and 17.
- Claims: 1-4, 6-17, 28-53 all partially same as invention 1 but for SEO 1D NO: 8, 7 and 18.
- Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 10, 9 and 19.
- Claims: 1-17, 20-53 all partially same as invention 1 but for SEO ID NO: 12, 11 and 20.
- Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 14, 13 and 21.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 16, 15 and 23.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 22.

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

- Claims: 1-17, 20-53 all partially
   same as invention i but for SEO ID NO: 42 and 41.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 48, 47 and 81.
- Claims: 1-4, 5-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 50, 49 and 82.
- 13. Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ 10 NO: 52 and 51.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 54, 53 and 83.
- 15. Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 56 and 55.
- 16. Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 58, 57 and 84.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 60, 59 and 85.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEQ ID NO: 62, 61 and 86.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 64, 63 and 79.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEQ ID NO: 66, 65 and 78.

## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEQ ID NO: 68 and 67.
- 22. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 70 and 69.
- 23. Claims: 1-4, 6-17, 28-53 all partially same as invention 1 but for SEQ ID NO: 72 and 71.
- Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEO ID NO: 75.
- 25. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEO ID NO: 76.
- 26. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEO 10 NO: 80.
- Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 88 and 87.
- 28. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 90 and 89.
- 29. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEO ID NO: 92 and 91.
- 30. Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 94 and 93.
- Claims: 1-4, 6-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 141, 140 and 169.

- Claims: 1-4, 6-17, 20-53 all partially
   same in invention 1 but for SEO ID NO: 143, 142 and 170.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO 1D NO: 145, 144 and 171.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEO ID NO: 147, 146 and 168.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEQ ID NO: 149, 148 and 73.
- Claims: 1-17, 20-53 all partially same as invention 1 but for SEQ ID NO: 151, 150 and 74.
- Claims: 1-4, 6-17, 20-53 all partially
   same as invention 1 but for SEQ 10 NO: 153, 152 and 77.
- 38. Claims: 11-17, 28-53 all partially, 18, 19

A fusion polypeptide comprising ESAT-6 or MPT59 each individually with one of the following epitope partners: DnaK, GroEL, urease, glutamine synthetase, the proline rich complex, L-alanine dehydrogenase, phosphate binding protein, 48 85 complex, HBMA, MPT51, MPT64, superoxide dismutase 19 kBa lipoprotein, alpha-crystallin, GroES, nucleic acids endoding said polypeptide, vaccines, pharmaceutical and immunological compositions containing said polypeptide or nucleic acid, an expression vector comprising said nucleic acid, a host transformed with said vector, immunization with said polypeptide, the use of said polypeptide in diagnosis, antibodies against said polypeptide.

information on patent family mambers

35 4500181 Application No PCT/DK 98/00132

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